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How useful is routine amniotic fluid and neonatal surface swab microbiology at Caesarean section?

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Abstract: BACKGROUND: Our aim was to evaluate the clinical impact of routine amniotic fluid and neonatal surface swab microbiology at Caesarean section. **MATERIALS AND METHODS:** Microbiology data from 1 537 neonates delivered by Caesarean section were analysed in the light of clinical outcome. **RESULTS:** 1 340 (87%) neonates had non-pathogenic bacteria or negative culture results from both amniotic fluid and surface swab samples. Of the 197 (13%) neonates with pathogenic bacteria, 22 (1.4%) were diagnosed with infection, but only in 6 (0.4%) were the bacteria presumed to be responsible for the infection. Amniotic fluid and surface swab culture had sensitivities of 54% and 35%, and positive predictive values of 14% and 17%, respectively, for detecting a neonate at risk of infection. **CONCLUSION:** Amniotic fluid and neonatal surface swab microbiology at Caesarean section contributes little if anything to postnatal management and can be safely dropped from operative routine.

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1 **How useful is routine amniotic fluid and neonatal surface swab microbiology at**
2 **caesarean section?**

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25 **Abstract**

26

27 **Background:** Our aim was to evaluate the clinical impact of routine amniotic fluid and
28 neonatal surface swab microbiology at caesarean section.

29 **Materials and methods:** Microbiology data from 1537 neonates delivered by
30 caesarean section were analysed in the light of clinical outcome.

31 **Results:** 1340 (87%) neonates had non-pathogenic bacteria or negative culture
32 results from both amniotic fluid and surface swab samples. Of the 197 (13%) neonates
33 with pathogenic bacteria, 22 (1.4%) were diagnosed with infection, but only in six (0.4%)
34 were the bacteria presumed responsible for the infection. Amniotic fluid and surface swab
35 culture had sensitivities of 54% and 35%, and positive predictive values of 14% and 17%,
36 respectively, for detecting a neonate at risk of infection.

37 **Conclusion:** Amniotic fluid and neonatal surface swab microbiology at caesarean
38 section contributes little if anything to postnatal management and can be safely dropped
39 from operative routine.

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43 **Key words:** amniotic fluid, surface swab, caesarean section, microbiology

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Introduction

Early detection of neonates at increased risk of infection is of major clinical interest.

Bacteriology of amniotic fluid samples and neonatal skin surface swabs at caesarean

section was proposed as a predictor of infection during the first days of life^{1, 2} and has

been routinely performed in some institutions. However, there is ongoing debate as to

whether isolates from these sources influence the development of neonatal infection and

subsequent clinical management. Some authors have attributed adverse perinatal

outcome to the bacteria isolated from amniotic fluid³⁻⁹ and have proposed sampling

amniotic fluid as an infection screening programme in preterms^{3, 4}. Others have

contended that bacterial invasion of the amniotic cavity does not increase the risk of

neonatal infection¹⁰⁻¹³. There is even debate over the effects of Ureaplasma urealyticum

on neonatal sepsis, meningitis and bronchopulmonary dysplasia¹⁴⁻¹⁶.

Studies on this issue are rare. Most were conducted decades ago^{1, 11} or limited to

subgroups such as preterm neonates or mothers with premature rupture of the

membranes^{6, 7, 9}. More particularly, sample sizes were small^{6, 11-13}.

Our aim was to evaluate the utility of routine amniotic fluid and neonatal surface swab

microbiology at caesarean section and its impact on subsequent clinical management,

regardless of gestational age or other limiting factors. A key purpose was to determine the

sensitivity and positive predictive value of the microbiology findings for neonatal infection.

65 **Material and Methods**

66 **Patient population.** In a retrospective study over 24 months (July 2003 – June 2005)
67 we analysed the microbiology data of all 1719 neonates delivered via caesarean section at
68 the Department of Obstetrics, University Hospital Zurich, Switzerland. We excluded 182
69 neonates on whom no amniotic fluid and skin surface microbiology had been performed.
70 Gestational age in the remaining 1537 neonates ranged from 24 to 43 weeks (median 38
71 weeks). Median birth weight was 2890g (range 260g to 5000g).

72 **Microbiological analysis and definitions.** Amniotic fluid samples (n=1321) and
73 neonatal cranial skin surface swabs (n=1486) were obtained at caesarean section.
74 Amniotic fluid samples were transported (Portagerm[®], bioMérieux, Marcy-l'Etoile, France)
75 to the microbiology laboratory for immediate Gram staining, aerobic culture on Columbia
76 sheep blood agar and chocolate agar, and anaerobic culture on Brucella agar (Becton,
77 Dickinson & Company, Franklin Lakes, NJ, (BD)) enriched with thioglycolate broth (BD).
78 A7 agar medium (bioMérieux) and Urée-Arginine Lyo 2[®] (bioMérieux), a ready-to-use
79 urea- and arginine-containing broth-based system for detecting urogenital mycoplasmas,
80 were used to detect Mycoplasma hominis and U. urealyticum. Surface swab samples were
81 cultured aerobically on sheep blood agar, McConkey agar, colistin-nalidixic acid agar,
82 chocolate agar, and streptococcal selective agar. Isolates were identified using standard
83 procedures. Microbiology data from the amniotic fluid samples, surface swabs and follow-
84 up samples (blood culture, cerebrospinal fluid and tracheal aspirate) were obtained from
85 the Institute of Medical Microbiology, University of Zurich.

86 Culture results were divided into two broad groups with respect to the clinical context:
87 pathogenic (e.g., Enterobacteriaceae, U. urealyticum and β -haemolytic Streptococcus
88 group B, in any amount) and non-pathogenic (e.g., lactobacilli, coagulase-negative
89 staphylococci and viridans streptococci, also in any amount). Classification of low-virulent

bacteria, i.e., mixed anaerobic bacteria, enterococci or peptostreptococci, depended on the amount present: low amounts or bacteria detected only on enrichment culture were classified as non-pathogenic; moderate or abundant amounts were classified as pathogenic. Negative cultures were pooled with the non-pathogenic results.

Clinical characteristics and definitions. Neonates were allocated to the following three groups: 1. No infection (no evidence of infection in the first six days of life); 2. Prophylactic antibiotics (administered over several days postpartum due to perinatal risk factors, e.g., mother positive for β -haemolytic Streptococcus group B, prolonged premature rupture of the membranes for >24 hours, and acute chorioamnionitis); 3. Infection (documented or suspected infection in the first six days of life). Clinical evidence of infection included respiratory distress syndrome, fever, hypotension, prolonged capillary refill time, hypoglycaemia and acidosis. Sepsis was diagnosed on the basis of a positive blood or cerebrospinal fluid culture combined with clinical signs. Clinical information was obtained from Zurich University Hospital's neonatal clinical database and individual patient records.

Statistical analysis. The sensitivity, specificity and positive and negative predictive value of a pathogenic culture result for detecting a neonate at risk of infection were separately calculated from cross-tabulations of the amniotic fluid and surface swab data. Pathogenic culture results were compared with non-pathogenic culture results. Neonates allocated to groups 2 & 3 (Prophylactic antibiotics & Infection) were considered together as at risk for infection and compared with those in group 1 (No infection).

111 Results

112 **Neonate characteristics.** Most neonates (1340/1537) had non-pathogenic or
 113 negative culture results from both amniotic fluid and surface swab (Figure 1). 1319/1340
 114 neonates showed no signs of infection (group 1); six neonates received prophylactic
 115 antibiotics (group 2) and 15 developed an infection (group 3). Cultures of amniotic fluid,
 116 surface swab or both were pathogenic in 197/1537 neonates, 170 of whom belonged to
 117 group 1, five to group 2, and the remaining 22 to group 3 (Infection).

118 **Analysis of the 22 neonates with infection and pathogenic amniotic fluid and/or**
 119 **surface swab cultures.** Microbiological workup was performed in 20/22 neonates, in 14 of
 120 whom additional cultures were negative or non-pathogenic. However, in the remaining six
 121 neonates, cultures were positive for pathogens: β -haemolytic Streptococcus group B
 122 (n=1), Klebsiella oxytoca (n=1), Escherichia coli (n=2) and other bacteria (n=2); each
 123 isolate was identical to that cultured from the amniotic fluid or surface swab, thus
 124 presumptive of a causal relationship with the neonatal infection.

125 **Microbiological analysis.** In total, 1321 amniotic fluid samples and 1486 surface
 126 swabs were tested; 430 and 456, respectively, proved positive, in most cases for more
 127 than one microorganism. In amniotic fluid, the most frequent isolates were coagulase-
 128 negative staphylococci (n=180), U. urealyticum (n=100) and Propionibacterium spp.
 129 (n=43). Pathogenic isolates comprised β -haemolytic Streptococcus group B (n=20), M.
 130 hominis (n=15), Klebsiella spp. (n=5) and E. coli (n=4). Surface swabs grew skin flora
 131 (n=83), enterococci (n=43), coagulase-negative staphylococci (n=38) and viridans
 132 streptococci (n=31); well-known pathogens were β -haemolytic Streptococcus group B
 133 (n=34) and E. coli (n=14).

134 **Amniotic fluid profile.** 1167/1321 amniotic fluid cultures were non-pathogenic.
 135 1149/1167 of these neonates belonged in group 1, six in group 2 and 12 in group 3.

136 Cultures of the remaining 154/1321 amniotic fluid samples grew pathogens; the neonates
137 concerned were distributed as follows: group 1, n=133; group 2, n=4; and group 3, n=17.
138 Pathogenic amniotic fluid culture had a sensitivity of 54% (21/39), a specificity of 90%
139 (1149/1282), a positive predictive value of 14% (21/154) and a negative predictive value of
140 99% (1149/1167) for detecting the risk of neonatal infection.

141 **Surface swab profile.** 1399/1486 surface swab cultures were non-pathogenic.
142 1371/1399 neonates belonged in group 1, nine in group 2 and 19 in group 3. Cultures of
143 the remaining 87/1486 swabs grew pathogens, with the neonates concerned distributed as
144 follows: group 1, n=72; group 2, n=1; and group 3, n=14. Pathogenic skin swab culture
145 had a sensitivity of 35% (15/43), a specificity of 95% (1371/1443), a positive predictive
146 value of 17% (15/87) and a negative predictive value of 98% (1371/1399) for detecting the
147 risk of neonatal infection.

148 Discussion

149 We evaluated the clinical impact of routine amniotic fluid and neonatal surface swab
150 microbiology at caesarean section regardless of specific clinical constellation. Pathogens
151 were detected in 197 (13%) of neonates, of whom only 22 (1.4%) developed an infection.
152 To test for a causal relationship between the amniotic fluid and/or surface swab pathogen
153 and the infection, we analysed the postnatal microbiology data and discovered that only in
154 six cases were the resulting isolates identical to those grown from the amniotic fluid or
155 surface swab. Thus pathogens detected at caesarean section can be presumed to have
156 accounted for postnatal infection in no more than 0.4% of the total 1537 cases studied.

157 The detection of infection risk by culturing amniotic fluid and neonatal surface swabs
158 had a sensitivity of only 54% and 35%, respectively. Sensitivities would have been even
159 lower if we had not considered neonates receiving prophylactic antibiotics (group 2) at risk
160 for infection. Moreover, the positive predictive values of 14% for amniotic fluid and 17% for
161 surface swabs reveal a disconnect between pathogen detection and development of
162 infection. This is consistent with reports of possible microbial invasion of the amniotic
163 cavity without demonstrable clinical signs of neonatal infection ¹². Conversely, non-
164 pathogenic cultures had high negative predictive values for infection: 99% for amniotic fluid
165 and 98% for surface swabs.

166 Not only does routine amniotic fluid and surface swab screening have a low risk
167 detection rate, it also provides no clinical information relevant to neonatal management. All
168 22 neonates identified with infection and amniotic fluid or surface swab pathogens had
169 already been treated with antibiotics due to their clinical presentation and risk factors. In
170 none of the 22 infected infants did the amniotic fluid or surface swab result influence
171 monitoring, antibiotic initiation or antibiotic choice.

172 Our data show that routine bacteriology of amniotic fluid and the neonatal surface at
173 caesarean section contributes little if anything to neonatal management. In our population,
174 most of the neonates were delivered at term. However, for preterms or neonates with
175 serious perinatal risk factors, amniotic fluid analysis might be useful to complement clinical
176 examination and microbiological workup; its positive predictive value might improve in this
177 setting. Skin swab analysis, on the other hand, has no value and should be discarded ⁴.

178

179 **Conclusion**

180 Routine amniotic fluid and neonatal surface swab bacteriology at caesarean section
181 contributes little if anything to clinical management. In view of its financial implications,
182 such screening should not be performed routinely.

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